International application No. PCT/AU03/01296

				
A.	CLASSIFICATION OF SUBJECT MA	ATTER		
Int. Cl. 7:	A61K 31/12, 31/352, 31/353, 31/555,	, 33/24,	A61P 35/00, C07F 15/00	
According to	International Patent Classification (IPC) o	r to both	national classification and IPC	·
В.	FIELDS SEARCHED			
Minimum docu	amentation searched (classification system follo	wed by o	lassification symbols)	
Documentation	searched other than minimum documentation	to the ext	ent that such documents are included in the fields search	ched
STN On-line	base consulted during the international search E Files MEDLINE, CA, WPIDS, BIOS histein, daidzein, flavan?, flavon?, plat	IS Key	data base and, where practicable, search terms used) words# dehydrocquol, platin?, cisplatin?, car plat? STN Substructure Search	boplatin?,
C.	DOCUMENTS CONSIDERED TO BE RE	LEVANI	r	
Category*	Citation of document, with indication, w	here app	жоргіate, of the relevant passages	Relevant to claim No.
		c Neuro	er 2001088116 surgery (2000 September), 33 (3), 123-31 n on growth inhibition and cytotoxicity of	·
X	human medulloblastoma cells".	Cispian	i on growth minoriton and cytoloxiony of	1-15
х	Biology, (1997 July) 17 (1) 51-9, "Ir	ican Jou nt er actio	er 97367506 urnal of Respiratory Cell and Molecular on with type II estrogen binding sites and ercetin in human non-small-cell lung	1-12, 14, 15
X F	urther documents are listed in the cont	inuation	n of Box C X See patent family ann	ex
"A" docume which is relevance "E" earlier a	categories of cited documents: ent defining the general state of the art is not considered to be of particular ce application or patent but published on or international filing date	ar o "X" d cr	nter document published after the international filing dained not in conflict with the application but cited to under theory underlying the invention ocument of particular relevance; the claimed invention onsidered novel or cannot be considered to involve anthen the document is taken alone	rstand the principle
claim(s) publicat reason (nt which may throw doubts on priority or which is cited to establish the tion date of another citation or other special as specified) nt referring to an oral disclosure, use,	"Y" d c: w a	ocument of particular relevance; the claimed invention onsidered to involve an inventive step when the document of or more other such documents, such combinated person skilled in the art ocument member of the same patent family	ent is combined
exhibition "P" docume	on or other means nt published prior to the international filing later than the priority date claimed		. ,	
Date of the actu 27 October 2	al completion of the international search		Date of mailing of the international search report - 7 NOV 2003	
AUSTRALIAN PO BOX 200, V	ng address of the ISA/AU PATENT OFFICE WODEN ACT 2606, AUSTRALIA pct@ipaustralia.gov.au (02) 6285 3929		Authorized officer S.R. IDRUS Telephone No: (02) 6283 2659	-

International application No.

C (Continua	tion). DOCUMENTS CONSIDERED TO BE RELEVANT	
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
P, X	STN File Medline, abstract accession number 2003459262 & TAMURA S. et. al., Pigment Cell Research, (2003 Oct) 16 (5) 470-6 "Genistein enhances the ciplatin-induced inhibitions of cell growth and apoptosis in human malignant melanoma cells".	1-15
	STN File Biosis, abstract accession number 2003:368481 & MANSOUR A. et. al., Blood (November 16, 2002) Vol.100, No. 11, pp. Abstract No. 4997	
P, X	"Enhancement of chemotherapeutic efficacy by combining agents that block IL-10 in CIL cell lines"	1-15
x	STN File Medline, abstract accession number 2002151803 & S. GIACOMELLI et. al., Life Sciences (8 February 2002), 70(12), pages 1447-1459, "Silybin and its bioavailable phospholipid complex (IdB 1016) potentiate in vitro and in vivo in the activity of cisplatin".	16-19
x	STN File Medline, abstract accession number 2000404390 & CALTAGIRONE S. et. al., International Journal of Cancer (15 August 2000), 87(4), pages 595-600, "Flavonoids apigenin and quercetin inhibit melanoma growth and metastatic potential".	16-19
x .	STN File Medline, abstract accession number 96273517 & SCAMBIA G. et. al., European Journal of Cancer (May 1996), 32A(5), pages 877-882, "Antiproliferative effect of silybin on gynaecological malignancies: synergism with cisplatin and doxorubicin".	16-19
x	STN File Medline, abstract accession number 91191699 & WAUD W. R. et. al., Cancer Chemotherapy and Pharmacology (1991), 27(6), pages 456-463, "Antitumor drug cross-resistance in vivo in a cisplatin-resistant murine P388 leukemia".	16-19
x	STN File Medline, abstract accession number 91107211 & NEELAM S. S. et. al., Investigational New Drugs (August 1990), 8(3), pages 263-268, "Combination of flavone acetic acid (FAA) with adriamycin, cisplatinum and difluoromethylornithine (DFMO) in vitro against human colon cancer cells".	16-19
x	STN File Medline, abstract accession number 92032982 & SCAMBIA G. et. al., Anti-Cancer Drugs (October 1990), 1(1), pages 45-48, "Synergistic antiproliferative activity of quercetin and cisplatin on ovarian cancer cell growth".	16-19
X	STN File CA, abstract number 119:131038 & DE VINCENZO R. et. al, Acta Medica Romana (1992), 30(1-2), pages 126-132, "Flavonoids and negative control of cell proliferation in ovarian tumors".	16-19
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International application No.
PCT/AU03/01296

	PC1/AU05/0	
C (Continuat	ion) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	ZYNER E. et. al., Acta Poloniae Pharmaceutica Drug Research, 1999, 56(2), pages 159-167, "Platinum (II) and palladium (II) N, O-chelates with substituted flavanone containing ligands".	
X	(see the whole document, in particular page 162, scheme 1 and page 164, table 1)	16-19
x	ZYNER E. et. al., Pharmazie, 1999, 54 (12), pages 945-946, "Pt(II) and Pd(II) complexes of 3-aminoflavone: in vitro and in vivo evaluation". (see the whole document)	16-19
x	LEI W. et. al., Anticancer Research, 1999, volume 19, pages 221-228, "Enhancement of chemosensitivity and programmed cell death by tyrosine kinase inhibitors correlates with EGFR expression in non-small cell lung cancer cells". (see the whole document, in particular the abstract and page 224)	16-19
	KANG B-J. et. al., Natural Product Sciences, 2000, 6(4), pages 165-169, "Scientific analysis of formulation theory of Chungpesagan-tang; in vitro cytotoxicity of cisplatin combined with Chungpesagan-tang".	16.10
X	(see the whole document, in particular the abstract and page 168)	16-19
x	WO 02/02548 A1 (ORION CORPORATION) 10 January 2002 See pages 2 and 7 and claims	1-15
x	WO 00/66576 A1 (G.J. CONSULTANTS PTY LTD) 9 November 2000 See pages 4-7, 12-20, examples 1, 2, 6-11	1-15
х	WO 99/49862 A1 (THE UNIVERSITY OF MISSISSIPPI) 7 October 1999 See page 2 and claim 1	1-15
x	WO 98/17662 A1 (NOVARTIS AG) 30 April 1998 See pages 2-3, 9-11, 13, 16, 17, examples 5, 9(d) and claims	1-15
x	EP 267155 A2 (ZYMA SA) 11 May 1988 See pages 2-8, formulas (I), (IIa), (IIb), (IIc), examples and claims	1-15
x	WO 80/02098 A1 (Z-L LIMITED PARTNERSHIP) 16 October 1980 See pages 2-8, 12-16, 18 20-23, claims	1-15
x	SEPULVEDA-BOZA, S et al, "The Preparation of New Isoflavones", Synthetic Communications (2001) vol 31 no 12 pages 1933-1940 See page 1935 compound 4f, page 1931 paragraph 1	1-15
	·	

International application No.

	FCT/A005/0	
C (Continua	tion) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	O'NEILL, M J et al, "Inducible Isoflavonoids from the Lima Bean, Phascolus lunatus",	
37	Phytochemistry (1986) vol 25 no 6 pages 1315-1322	
X	See page 1316 column 1 compound 19 and column 2 paragraph 2	1-15
	WOLFBEIS, O S et al, "The Absorption and Fluorescence of Isoflavones and the	
	Effect of Shift Reagents", Z. Naturforsch. (1984) 39b pages 238-243	
X	See page 240 Table 1 compounds 2, 8, 11, 13, 15, 18	1-15
	ARORA, S K et al, "The Synthesis of Tlatlancuayin", Tetrahedron (1962) vol 18 pages	·
	559-565 See page 559, page 560 compounds (V)-(VII) and paragraph 3, page 564 paragraphs 5	
X	and 6, page 565	1-15
	STN Chemical Abstract Accession No 135:355315	
X	& Chemical And Pharmaceutical Bulletin (2001), 49 (9), 1229-1231	1-15
	STN Chemical Abstract Accession No 135:121648	
X	& Journal of Agricultural and Food Chemistry (2001), 49 (6) 3024-3033	1-15
77	STN Chemical Abstract Accession No 124:341448	
X	& Archives of Microbiology (1995), 164 (6), 428-34	1-15
	STN Chemical Abstract Accession No 124:316797	
X	& Chemical And Pharmaceutical Bulletin (1996), 44 (3), 486-91	1-15
	STN Chemical Abstract Accession No 124:140985	
·X	& Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1995), 37th, 493-8	1-15
	STN Chemical Abstract Accession No 115:68424	
X	& Phytochemistry (1991), 30 (4), 1281-4	1-15
3 L ,	C. I hy toolicanistry (1991), 50 (4), 1201-4	. 1-15
	STN Chemical Abstract Accession No 114:41246	
X	& Angewandte Botanik (1990), 64 (1-2), 175-90	1-15
	STN Chemical Abstract Accession No 112:69573	
X	& International Journal of Tissue Reactions (1989), 11 (3), 107-12	1-15
	STN Chemical Abstract Accession No 102:59329	
X	& Phytochemistry (Elsevier) (1984), 23 (11), 2703-4	1-15
	(11), 2703	1-13
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International application No. PCT/AU03/01296

C (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT			
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
	STN Chemical Abstract Accession No 102:59220		
X	& Phytochemistry (Elsevier) (1984), 23 (6), 1342-3	1-15	
	STN Chemical Abstract Accession No 97:109739		
X	& J. Chem. Soc., Perkin Trans. 1 (1982), (6), 1389-94	1-15	
	STN Chemical Abstract Accession No 95:111690		
X	& Phytochemistry (1981), 20 (4), 799-801	1-15	
	STN Chemical Abstract Accession No 82:97918		
X	& J. Inst. Chem., Calcutta (1974), 46, Pt. 3, 61-5	1-15	
	STN Chemical Abstract Accession No 76:140428		
X	& J. Inst. Chem., Calcutta (1971), 43 (6), 234-40	1-15	
	STN Chemical Abstract Accession No 70:57577		
X	Indian J. Chem. (1968), 6 (9), 481-4	1-15	
	STN Chemical Abstract Accession No 63:54537		
X	& Bull. Chem. Soc. Japan (1965), 38 (6), 887-93	1-15	
	STN Chemical Abstract Accession No 61:61569		
X	& Periodica Polytech. (1963), 7 (4), 241-58	1-15	
	STN Chemical Abstract Accession No 126:139728		
	(see CAS RN 116718-84-4)		
X	& Atherosclerosis (1997), 128(1), 59-66	1-15	
	JHA, H C et al, "Carbon-13 Chemical Shift Assignments of Chromones and	}	
	Isoflavones", Can. J. Chem. (1980) vol 58 no 12 pages 1211-1219		
X	See pages 1212-1213 Table 1(b) compounds 15, 17, 34-39, 41, 49-50	1-15	
	STN Chemical Abstracts Accession No 128:164027		
х	& Antioxidants in Health and Disease (1998), 7 (Flavonoids in Health and Disease pages 295-302	e) 1-15	
7.	, here and a second sec	'	
	STN Chemical Abstracts Accession No 117:124019		
X	& Biochemical Pharmacology (1992) vol 44(1), pages 157-162	1-15	
	·		

International application No.

Box I	Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This internative	ional search report has not been established in respect of certain claims under Article 17(2)(a) for the following
1.	Clairns Nos :
	because they relate to subject matter not required to be searched by this Authority, namely:
2. X	<u> </u>
	because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
	(i) as claims 1-15 are broadly drafted it is not economically possible to search the full scope of the claims. With due consideration to the examples, the search was limited to the isoflavones, and
	(ii) claims 16-19 are directed to platinum-isoflavanoid complexes of formula (II), wherein the isoflavanoid ligand is as exemplified in the specification (compounds 1-30 on pages 19-22);
3.	Claims Nos:
	because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a)
Вох П	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This Internat	onal Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on I	rotest The additional search fees were accompanied by the applicant's protest.
٠	No protest accompanied the payment of additional search fees.

Information on patent family members

International application No.

PCT/AU03/01296

This Annex lists the known "A" publication level patent family members relating to the patent documents cited in the above-mentioned international search report. The Australian Patent Office is in no way liable for these particulars which are merely given for the purpose of information.

Paten	t Document Cited in Search Report			Pate	ent Family Member		
wo	9949862	AU	34564/99				
wo	9817662	AU	49479/97	•			
EP	267155	AU	80655/87	DD	275048	DK	5756/87
		FI	874804	HU	48611	JР	63130589
		NO	874489	NZ	222411	PT	86055
		US	4814346	ZA	8708245		
wo	8002098	US	4264509	BR	7909002	CA	1140560
		DK	5288/80	EP	27796	NL	7906193
		US	4366082	US	4366248	US	4390559
		US	4157984	· US	4234577	US	4368264
		US	4218489	US	4232122	BR	7908996
		DK	4928/80	EP	25783	NL	7906287
		wo	8002027				•
wo	0202548	AU	72597/01	FI	20001593		
wo	0066576	EP	1189897				•
				•			END OF ANNEX

PATENT COOPERATION TREATY PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference 78273853		tification of Transmittal of International Search Report PCT/ISA/220) as well as, where applicable, item 5 below.
International application No.	International filing date (day/mont	
PCT/AU03/01296	2 October 2003	2 October 2002
Applicant NOVOGEN RESEARCH PTY	LTD et al	
This international search report has been preparticle 18. A copy is being transmitted to the		Authority and is transmitted to the applicant according to
This international search report consists of a	total of 8 sheets.	
It is also accompanied by a cop	y of each prior art document cited in	this report.
1. Basis of the report		
which it was filed, unless other	wise indicated under this item.	at on the basis of the international application in the language in
the international search w (Rule 23.1(b)).	as carried out on the basis of a trans	lation of the international application furnished to this Authority
b. With regard to any nucleotide carried out on the basis of the s		ed in the international application, the international search was
contained in the internation	onal application in written form.	·
filed together with the int	ernational application in computer r	eadable form.
furnished subsequently to	this Authority in written form.	
	this Authority in computer readable	
the statement that the sub application as filed has be		e listing does not go beyond the disclosure in the international
the statement that the info	ormation recorded in computer reada	ble form is identical to the written sequence listing has been
2. X Certain claims were found un	searchable (See Box I).	
3. Unity of invention is lacking (See Box II).	
4. With regard to the title,	the text is approved as submitted by	by the applicant.
	the text has been established by th	is Authority to read as follows:
5. With regard to the abstract,	the text is approved as submitted l	by the applicant
	the text has been established, according to the applicant may, within one mosubmit comments to this Authority	ording to Rule 38.2(b), by this Authority as it appears in Box III. onth from the date of mailing of this international search report, y.
6. The figure of the drawings to be pub	lished with the abstract is Figure No).
	as suggested by the applicant.	X None of the figures
	because the applicant failed to sup	ggest a figure
	because this figure better characte	erizes the invention



Box I Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)	
This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:	
1. Claims Nos:	
because they relate to subject matter not required to be searched by this Authority, namely:	
2. X Claims Nos: 1-15, 16-19	
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:	
(i) as claims 1-15 are broadly drafted it is not economically possible to search the full scope of the	
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3. Claims Nos:	
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a)	
Box II Observations where unity of invention is lacking (Continuation of item 3 of first sheet)	
This International Searching Authority found multiple inventions in this international application, as follows:	
·	
As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims	
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As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:	
No required additional search fees were timely paid by the applicant. Consequently, this international search repois restricted to the invention first mentioned in the claims; it is covered by claims Nos.:	rt
Remark on Protest The additional search fees were accompanied by the applicant's protest.	
No protest accompanied the payment of additional search fees.	

Α.	CLASSIFICATION OF SUBJECT MATTER		
	A61K 31/12, 31/352, 31/353, 31/555, 33/24, A		
According to I	international Patent Classification (IPC) or to both	national classification and IPC	
В.	FIELDS SEARCHED		
Minimum docu	mentation searched (classification system followed by classification system followed system follow	assification symbols)	
Documentation	searched other than minimum documentation to the exte	nt that such documents are included in the fields search	hed
STN On-line	base consulted during the international search (name of a Files MEDLINE, CA, WPIDS, BIOSIS Keywistein, daidzein, flavan?, flavon?, platin?, cisp	ords# dehydroequol, platin?, cisplatin?, carb	ooplatin?,
c.	DOCUMENTS CONSIDERED TO BE RELEVANT		
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х	STN File Medline, abstract accession number & KHOSHYOMN S. et al., Pediatric Neuros "Synergistic action of genistein and cisplatin human medulloblastoma cells".	surgery (2000 September), 33 (3), 123-31	1-15
х	STN File Medline, abstract accession number & CALTAGIRONE S. et. al., American Journ Biology, (1997 July) 17 (1) 51-9, "Interaction atiproliferative activity of tamoxifen and que cancer".	rnal of Respiratory Cell and Molecular n with type II estrogen binding sites and	1-12, 14, 15
X F	urther documents are listed in the continuation	of Box C X See patent family annual	ex
"A" docume which i relevand "E" earlier a	s not considered to be of particular ce application or patent but published on or "X" c international filing date cc	ter document published after the international filing dand not in conflict with the application but cited to under theory underlying the invention ocument of particular relevance; the claimed invention onsidered novel or cannot be considered to involve an	rstand the principle
"L" docume claim(s publica reason (ent which may throw doubts on priority "Y" do) or which is cited to establish the co tion date of another citation or other special w (as specified) a	hen the document is taken alone ocument of particular relevance; the claimed invention onsidered to involve an inventive step when the documith one or more other such documents, such combinative person skilled in the art ocument member of the same patent family	ent is combined
date bu	t later than the priority date claimed al completion of the international search	Date of mailing of the international search report	
27 October 2		- 7 NOV 2003	
AUSTRALIAN PO BOX 200, E-mail address	ing address of the ISA/AU I PATENT OFFICE WODEN ACT 2606, AUSTRALIA : pct@ipaustralia.gov.au (02) 6285 3929	Authorized officer S.R. IDRUS Telephone No: (02) 6283 2659	- .



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P, X	STN File Medline, abstract accession number 2003459262 & TAMURA S. et. al., Pigment Cell Research, (2003 Oct) 16 (5) 470-6 "Genistein enhances the ciplatin-induced inhibitions of cell growth and apoptosis in human malignant melanoma cells".	1-15
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x	STN File Medline, abstract accession number 2002151803 & S. GIACOMELLI et. al., Life Sciences (8 February 2002), 70(12), pages 1447-1459, "Silybin and its bioavailable phospholipid complex (IdB 1016) potentiate in vitro and in vivo in the activity of cisplatin".	16-19
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	159-167, "Platinum (II) and palladium (II) N, O-chelates with substituted flavanone	
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	ZYNER E. et. al., Pharmazie, 1999, 54 (12), pages 945-946, "Pt(II) and Pd(II)	
	complexes of 3-aminoflavone: in vitro and in vivo evaluation".	
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	with EGFR expression in non-small cell lung cancer cells".	
X	(see the whole document, in particular the abstract and page 224)	16-19
	KANG B-J. et. al., Natural Product Sciences, 2000, 6(4), pages 165-169, "Scientific	
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	combined with Chungpesagan-tang".	
X :	(see the whole document, in particular the abstract and page 168)	16-19
	WO 02/02548 A1 (ORION CORPORATION) 10 January 2002	
X	See pages 2 and 7 and claims	1-15
	WO ON COST A LOCAL CONICHI TANTE PTV I TOVO November 2000	
x	WO 00/66576 A1 (G.J. CONSULTANTS PTY LTD) 9 November 2000 See pages 4-7, 12-20, examples 1, 2, 6-11	1-15
^	See pages 4-7, 12-20, examples 1, 2, 0-11	
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X	See page 2 and claim 1	1-15
	WO 98/17662 A1 (NOVARTIS AG) 30 April 1998	
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	ED 267155 A2 (737MA CA) 11 May 1000	
x	EP 267155 A2 (ZYMA SA) 11 May 1988 See pages 2-8, formulas (I), (IIa), (IIb), (IIc), examples and claims	1-15
Λ	bee pages 2-0, formulas (1), (110), (110), (110), examples and elamine	
	WO 80/02098 A1 (Z-L LIMITED PARTNERSHIP) 16 October 1980	
X	See pages 2-8, 12-16, 18 20-23, claims	1-15
•	SEPULVEDA-BOZA, S et al, "The Preparation of New Isoflavones", Synthetic	
37	Communications (2001) vol 31 no 12 pages 1933-1940	1-15
X	See page 1935 compound 4f, page 1931 paragraph 1	1-13



C (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT					
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	WOLFBEIS, O S et al, "The Absorption and Fluorescence of Isoflavones and the				
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X	See page 240 Table 1 compounds 2, 8, 11, 13, 15, 18				
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